新**州** 网络克莱拉

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We claim:

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- 1. A soluble purified PHEX enzyme (secPHEX) and variants thereof, comprising a PHEX ectodomain or catalytic part thereof.
- 2. A soluble purified PHEX enzyme (secPHEX) and variants thereof, comprising a PHEX ectodomain or catalytic part thereof and a PHEX signal peptide/transmembrane region modified so as to confer solubility to said PHEX enzyme when expressed in a eukaryotic host, wherein said host is not a human being:
 - 3. A mutant of the enzyme of claim, 1 or 2, which is inactive but retains binding capacity to a ligand molecule to PHEX.
 - 4. A mutant as defined in claim 3 consisting of PHEX enzyme having the glutamic acid residue at position 581 is mutated.
 - 5. A mutant as defined in claim 3 consisting of PHEX enzyme having the glutamic acid residue at position 581 substituted with high a hydrophobic amino acid residue.
 - 6. A mutant as defined in claim 3 consisting of PHEX enzyme having the glutamic acid residue at position 581 substituted with a valine residue.
 - Annucleic acid which comprises a truncated PHEX gene sequence encoding PHEX membrane-anchor domain modified to include a cleavable signal peptide and PHEX C-terminal control ectodomain is active or inactive.
 - 8. A recombinant vector comprising the nucleic acid of claim 7.

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- 9. A recombinant vector as defined in claim 8, which is an expression vector.
- 10. A eukaryotic host comprising the recombinant vector of claim 8, wherein said host is not a human being
- 11. A eukaryotic host comprising the recombinant vector of claim 9, wherein said host is not a human being.
- 12. A method for producing a soluble PHEX enzyme or an inactive mutant thereof, which comprises the steps of:
 - allowing the eukaryotic host of claim .10 or 11 to express said nucleic acid, and
 - recovering the soluble PHEX enzyme or mutant thereof as a secretion product of said host.
- 13. An antigenic composition, which comprises the enzyme of any one of claims 1 to 6.
 - 14. An antibody capable of binding to PHEX and raised against the enzyme of any one of claims 1 to 6 or fragment thereof.
- The antibody of claim 14, wherein said fragment extends from residue 121 to residue 294 of the amino acid sequence of PHEX.
 - 16. An antibody as defined in claim 14, which is a monoclonal antibody.
- Aniantibody as defined in claim 15, which is a monoclonal antibody.

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- 18. The antibody of claim 16, which is a PHEX neutralizing antibody.
 - A hybridoma producing the antibody of any one of claims 16 to 19. 18.
 - A composition comprising the enzyme of claim 1 or 2 or the 20. nucleic acid of claim 7 and a pharmaceutically acceptable carrier. 副司部 田子
 - 21. A composition complising the enzyme of any one of claims 3 to 6 and a pharmaceutically acceptable carrier.
 - A composition comprising the antibody of any one of claims 14 to 22. 18 and a pharmaceutically acceptable carrier. 養養養養 海拔系统 俄山的一种工作
 - A diagnostic reagent for detecting the presence or amount of PHEX, comprising the antibody of any one of claims 14 to 18.
 - A diagnostic kit for detecting the presence or amount of PHEX 24. comprising the antibody of any one of claims 14 to 18. 福州学科强力工程
 - 25. The diagnostic kit of claim 24, which further comprises a soluble PHEX enzyme.
 - 26. A method for detecting the presence or an amount of PHEX in a sample, which comprises the steps of:
 - contacting said sample with the antibody of any of any one of claims 14 to 18 in conditions such that the immune complex can form; and 中疆。湖湖南区,现代中央旅游发展。
 - detecting the immune complexed as an indication of the presence or amount of PHEX in said sample.

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- A device for purifying PHEX or a mutant thereof which comprises 27. the antibody of any one of claims 14 to 18.
- A device for screening AHEX ligands, which comprises the soluble PHEX enzyme or a mutant thereof as defined in any one of claims 1 to 6. 7番 明诗 、明代县原奉。
- 29. The device of claim 27, wherein said antibody is fixed onto a · 建铁铁铁矿 (1) (1) (1) (1) solid support.
- fixed onto a solid support. 1、16種以關係、1個自動能量於如本
 - The device of claim 30, wherein said PHEX enzyme or mutant is 31. fixed onto the solid support through its binding to an anti-PHEX antibody itself fixed onto said solid support. 精練 建铁头 銀小兒 电工学
 - The device of claim 30, wherein said PHEX enzyme or mutant is 32. fixed onto the solid support through a C-terminal amino acid extension ending with a residue or group capable of coupling PHEX to the solid support.;

響電音 33. 報Armethod for obtaining a PHEX Ilgand which comprising the steps of:

> contacting a sample containing one or more molecules with a PHEX enzyme of mutant as defined in any one of claims 1 to 6 in conditions such that binding of said one or more molecules with PHEX can occur:

a PHEX ligand in said sample; and

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| 接触性,性量中心selecting said\PHEX ligand.

- 34. The method of claim 33, wherein said ligand is a PHEX/inhibitor with or substrate.
- 35. A method for evaluating the activity of a molecule for its capacity of heing a substrate of PHEX comprising the steps of:
 - contacting said molecule with the PHEX enzyme of claim.
 1 in substantially phosphate-free conditions; and
 observing a cleavage product of said molecule as an indication that the molecule is a PHEX substrate.
 - 38. The method of claim 35, which further comprises the step of comparing said molecule with PTHrP107-139 as a positive control.
 - A method for evaluating PHEX activity in a sample which comprises the steps of contacting the sample with a substrate as defined in claim 35 or 36, or preferably with PTHrP107-139, in substantially phosphate-free conditions and observing the apparition of a cleavage product of said substrate or PTHrP107-139 as an indication of PHEX activity in the sample.
- 38. The method of claim 37, which further comprises a step of comparing said PHEX activity in the sample with the activity of the PHEX enzyme of claim 1 as a positive control.
 - 39. A method for evaluating the activity of a molecule for its capacity of being an inhibitor of PHEX comprising the steps of:

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阿拉斯特特的 医中心性神经病

"福州县建筑"通过1200年,1900年,1900年

经销售的 医红色性 医二十二

"特殊"的一种,

"曾华德姓",然后这种种事的多少多

化工作 计差别操业分析 "好好好"

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contacting said molecule with a substrate as defined in claim 35 or 36, or preferably with PTHrP107-139, and the PHEX enzyme of claim 1 in substantially phosphate-free conditions; and observing an inhibition of the formation of a cleavage product as an indication that said molecule is a PHEX inhibitor.

40. A kit for executing the method of any one of claims 35 to 39.

1、 海洲 印源中省核社会

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